

**CLAIMS**

1. A polypeptide, which polypeptide:

- (i) comprises the amino acid sequence as recited in SEQ ID NO:6 and/or SEQ ID NO:8;
- 5 (ii) is a fragment thereof which functions as a member of the defensin family of proteins, or having an antigenic determinant in common with the polypeptide of (i); or
- (iii) is a functional equivalent of (i) or (ii).

2. A polypeptide according to claim 1 which consists of the amino acid sequence as recited  
10 in SEQ ID NO:6 and/or SEQ ID NO:8.

3. A polypeptide, which polypeptide:

- (i) comprises the amino acid sequence as recited in SEQ ID NO:14 and/or SEQ ID NO:16;
- 15 (ii) is a fragment thereof which functions as a member of the defensin family of proteins, or having an antigenic determinant in common with the polypeptide of (i); or
- (iii) is a functional equivalent of (i) or (ii).

4. A polypeptide according to claim 3 which consists of the amino acid sequence as recited  
in SEQ ID NO:14 and/or SEQ ID NO:16.

20 5. A polypeptide which is a functional equivalent according to part (iii) of claim 1 or claim 3, characterised in that it is homologous to the amino acid sequence as recited in SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:14 or SEQ ID NO:16 and is a member of the defensin family of proteins.

25 6. A polypeptide which is a fragment or a functional equivalent as recited in any one of claims 1, 3 or 5, which has greater than 80% sequence identity with the amino acid sequence recited in SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:14 or SEQ ID NO:16 or with an active fragment thereof, preferably greater than 85%, 90%, 95%, 98% or 99% sequence identity.

7. A polypeptide which is a functional equivalent as recited in any one of claims 1, 3, 5 or 6, which exhibits significant structural homology with a polypeptide having the amino acid sequence recited SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:14 or SEQ ID NO:16.
8. A polypeptide which is a fragment as recited in any one of claims 1, 3 and claim 6  
5 having an antigenic determinant in common with the polypeptide of part (i) of claim 1 or claim 3 which consists of 7 or more amino acid residues from the amino acid sequence recited in SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:14 or SEQ ID NO:16 .
9. A purified nucleic acid molecule which encodes a polypeptide according to any one of the preceding claims.
- 10 10. A purified nucleic acid molecule according to claim 9, which comprises the nucleic acid sequence as recited in SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:13 and/or SEQ ID NO:15, or is a redundant equivalent or fragment thereof.
11. A purified nucleic acid molecule according to claim 9 or claim 10 which consists of the nucleic acid sequence as recited in SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:13 and/or SEQ ID NO:15, or is a redundant equivalent or fragment thereof.  
15
12. A purified nucleic acid molecule which hybridizes under high stringency conditions with a nucleic acid molecule according to any one of claims 9 to 11.
13. A vector comprising a nucleic acid molecule as recited in any one of claims 9 to 12.
14. A host cell transformed with a vector according to claim 13.
- 20 15. A ligand which binds specifically to the defensin polypeptide according to any one of claims 1 to 8.
16. A ligand according to claim 15, which is an antibody.
17. A compound that either increases or decreases the level of expression or activity of a polypeptide according to any one of claims 1 to 8.  
25
18. A compound according to claim 17 that binds to a polypeptide according to any one of claims 1 to 8 without inducing any of the biological effects of the polypeptide.
19. A compound according to claim 18, which is a natural or modified substrate, ligand, enzyme, receptor or structural or functional mimetic.
20. A polypeptide according to any one of claims 1 to 8, a nucleic acid molecule according

to any one of claims 9 to 12, a vector according to claim 13, a host cell according to claim 14, a ligand according to claim 15 or claim 16, or a compound according to any one of claims 17 to 19, for use in therapy or diagnosis of disease.

21. A method of diagnosing a disease in a patient, comprising assessing the level of expression of a natural gene encoding a polypeptide according to any one of claims 1 to 8, or assessing the activity of a polypeptide according to any one of claims 1 to 8, in tissue from said patient and comparing said level of expression or activity to a control level, wherein a level that is different to said control level is indicative of disease.  
5
22. A method according to claim 21 that is carried out *in vitro*.
- 10 23. A method according to claim 21 or claim 22, which comprises the steps of: (a) contacting a ligand according to claim 15 or claim 16 with a biological sample under conditions suitable for the formation of a ligand-polypeptide complex; and (b) detecting said complex.
24. A method according to claim 21 or claim 22, comprising the steps of:  
15     a) contacting a sample of tissue from the patient with a nucleic acid probe under stringent conditions that allow the formation of a hybrid complex between a nucleic acid molecule according to any one of claims 9 to 12 and the probe;  
         b) contacting a control sample with said probe under the same conditions used in step a); and  
20         c) detecting the presence of hybrid complexes in said samples; wherein detection of levels of the hybrid complex in the patient sample that differ from levels of the hybrid complex in the control sample is indicative of disease.
25. A method according to claim 21 or claim 22, comprising:  
         a) contacting a sample of nucleic acid from tissue of the patient with a nucleic acid primer under stringent conditions that allow the formation of a hybrid complex between a nucleic acid molecule according to any one of claims 9 to 12 and the primer;  
         b) contacting a control sample with said primer under the same conditions used in step a); and  
         c) amplifying the sampled nucleic acid; and  
30         d) detecting the level of amplified nucleic acid from both patient and control samples;

wherein detection of levels of the amplified nucleic acid in the patient sample that differ significantly from levels of the amplified nucleic acid in the control sample is indicative of disease.

26. A method according to claim 21 or claim 22 comprising:

- 5      a)obtaining a tissue sample from a patient being tested for disease;
- b)isolating a nucleic acid molecule according to any one of claims 9 to 12 from said tissue sample; and
- c)diagnosing the patient for disease by detecting the presence of a mutation which is associated with disease in the nucleic acid molecule as an indication of the disease.

10 27. The method of claim 26, further comprising amplifying the nucleic acid molecule to form an amplified product and detecting the presence or absence of a mutation in the amplified product.

28. The method of claim 26 or claim 27, wherein the presence or absence of the mutation in the patient is detected by contacting said nucleic acid molecule with a nucleic acid probe that hybridises to said nucleic acid molecule under stringent conditions to form a hybrid double-stranded molecule, the hybrid double-stranded molecule having an unhybridised portion of the nucleic acid probe strand at any portion corresponding to a mutation associated with disease; and detecting the presence or absence of an unhybridised portion of the probe strand as an indication of the presence or absence of a disease-associated mutation.

15 29. A method according to any one of claims 21 to 28, wherein said disease includes, but is not limited to, cell proliferative disorders, including neoplasm, melanoma, lung, colorectal, breast, pancreas, head and neck and other solid tumours; myeloproliferative disorders, such as leukemia, non-Hodgkin lymphoma, leukopenia, thrombocytopenia, angiogenesis disorder, Kaposi's sarcoma; autoimmune/inflammatory disorders, including allergy, inflammatory bowel disease, arthritis, psoriasis and respiratory tract inflammation, asthma, and organ transplant rejection; cardiovascular disorders, including hypertension, oedema, angina, atherosclerosis, thrombosis, sepsis, shock, reperfusion injury, and ischemia; neurological disorders including central nervous system disease, Alzheimer's disease, brain injury, amyotrophic lateral sclerosis, and pain; developmental disorders; metabolic disorders including diabetes mellitus,

20

25

30

osteoporosis, and obesity, AIDS and renal disease; infections including viral infection, bacterial infection, fungal infection, parasitic infection, sublethal endotoxaemia, septic shock, microbial infection of the amniotic cavity, Jarish-Herxheimer reaction of relapsing fever, infectious diseases of the central nervous system, acute pancreatitis, 5 ulcerative colitis, empyaema, haemolytic uraemic syndrome, meningococcal disease, gastric infection, pertussis, peritonitis, psoriasis, rheumatoid arthritis, sepsis, asthma, HIV and glomerulonephritis.

30. A method according to any one of claims 21 to 28, wherein said disease is a disease in which defensins are implicated.

10 31. Use of a polypeptide according to any one of claims 1 to 8 as a defensin protein.

32. A pharmaceutical composition comprising a polypeptide according to any one of claims 1 to 8, a nucleic acid molecule according to any one of claims 9 to 12, a vector according to claim 13, a host cell according to claim 14, a ligand according to claim 15 or claim 16, or a compound according to any one of claims 17 to 19.

15 33. A vaccine composition comprising a polypeptide according to any one of claims 1 to 8 or a nucleic acid molecule according to any one of claims 9 to 12.

34. A polypeptide according to any one of claims 1 to 8, a nucleic acid molecule according to any one of claims 9 to 12, a vector according to claim 13, a host cell according to claim 14, a ligand according to claim 15 or claim 16, a compound according to any one of claims 17 to 19, or a pharmaceutical composition according to claim 32, for use in 20 the manufacture of a medicament for the treatment of cell proliferative disorders, including neoplasm, melanoma, lung, colorectal, breast, pancreas, head and neck and other solid tumours; myeloproliferative disorders, such as leukemia, non-Hodgkin lymphoma, leukopenia, thrombocytopenia, angiogenesis disorder, Kaposi's sarcoma; autoimmune/inflammatory disorders, including allergy, inflammatory bowel disease, 25 arthritis, psoriasis and respiratory tract inflammation, asthma, and organ transplant rejection; cardiovascular disorders, including hypertension, oedema, angina, atherosclerosis, thrombosis, sepsis, shock, reperfusion injury, and ischemia; neurological disorders including central nervous system disease, Alzheimer's disease, brain injury, amyotrophic lateral sclerosis, and pain; developmental disorders; 30 metabolic disorders including diabetes mellitus, osteoporosis, and obesity, AIDS and renal disease; infections including viral infection, bacterial infection, fungal infection,

parasitic infection, sublethal endotoxaemia, septic shock, microbial infection of the amniotic cavity, Jarish-Herxheimer reaction of relapsing fever, infectious diseases of the central nervous system, acute pancreatitis, ulcerative colitis, empyaema, haemolytic uraemic syndrome, meningococcal disease, gastric infection, pertussis, peritonitis, 5 psoriasis, rheumatoid arthritis, sepsis, asthma, HIV and glomerulonephritis.

35. A polypeptide according to any one of claims 1 to 8, a nucleic acid molecule according to any one of claims 9 to 12, a vector according to claim 13, a host cell according to claim 14, a ligand according to claim 15 or claim 16, a compound according to any one of claims 17 to 19, or a pharmaceutical composition according to claim 32, for use in 10 the manufacture of a medicament for the treatment of a disease in which defensins are implicated.
36. A method of treating a disease in a patient, comprising administering to the patient a polypeptide according to any one of claims 1 to 8, a nucleic acid molecule according to any one of claims 9 to 12, a vector according to claim 13, a host cell according to claim 14, a ligand according to claim 15 or claim 16, a compound according to any one of 15 claims 17 to 19, or a pharmaceutical composition according to claim 32.
37. A method according to claim 36, wherein, for diseases in which the expression of the natural gene or the activity of the polypeptide is lower in a diseased patient when compared to the level of expression or activity in a healthy patient, the polypeptide, 20 nucleic acid molecule, vector, ligand, compound or composition administered to the patient is an agonist.
38. A method according to claim 36, wherein, for diseases in which the expression of the natural gene or activity of the polypeptide is higher in a diseased patient when compared to the level of expression or activity in a healthy patient, the polypeptide, nucleic acid molecule, vector, ligand, compound or composition administered to the 25 patient is an antagonist.
39. A method of monitoring the therapeutic treatment of disease in a patient, comprising monitoring over a period of time the level of expression or activity of a polypeptide according to any one of claims 1 to 8, or the level of expression of a nucleic acid molecule according to any one of claims 9 to 12 in tissue from said patient, wherein altering said level of expression or activity over the period of time towards a control 30 level is indicative of regression of said disease.

40. A method for the identification of a compound that is effective in the treatment and/or diagnosis of disease, comprising contacting a polypeptide according to any one of claims 1 to 8, or a nucleic acid molecule according to any one of claims 9 to 12 with one or more compounds suspected of possessing binding affinity for said polypeptide or nucleic acid molecule, and selecting a compound that binds specifically to said nucleic acid molecule or polypeptide.  
5
41. A kit useful for diagnosing disease comprising a first container containing a nucleic acid probe that hybridises under stringent conditions with a nucleic acid molecule according to any one of claims 9 to 12; a second container containing primers useful  
10 for amplifying said nucleic acid molecule; and instructions for using the probe and primers for facilitating the diagnosis of disease.
42. The kit of claim 41, further comprising a third container holding an agent for digesting unhybridised RNA.
43. A kit comprising an array of nucleic acid molecules, at least one of which is a nucleic  
15 acid molecule according to any one of claims 9 to 12.
44. A kit comprising one or more antibodies that bind to a polypeptide as recited in any one of claims 1 to 8; and a reagent useful for the detection of a binding reaction between said antibody and said polypeptide.
45. A transgenic or knockout non-human animal that has been transformed to express  
20 higher, lower or absent levels of a polypeptide according to any one of claims 1 to 8.
46. A method for screening for a compound effective to treat disease, by contacting a non-human transgenic animal according to claim 45 with a candidate compound and determining the effect of the compound on the disease of the animal.